

In the Specification:

Please replace the paragraph beginning at page 2, line 32, with the following:

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--Fig. 2 is a chart showing the alignment of predicted Rbx1 protein sequences from human (SEQ ID NO:1), mouse (SEQ ID NO:1), *Drosophila melanogaster* (SEQ ID NO:6), *Caenorhabditis elegans* (SEQ ID NO:7), and *Saccharomyces cerevisiae* (SEQ ID NO:2) with APC11 from *S. cerevisiae* (SEQ ID NOS:8 and 9), wherein DROS = *Drosophila melanogaster*; ELEGANS = *Caenorhabditis elegans*; and YEAST = *Saccharomyces cerevisiae*. The alignment was generated using the MACAW program. (Schuler, *et al.* 1991. "A workbench for multiple alignment construction and analysis," *Proteins; Struct Funct Genet* 9:180-190). Dark shading indicates positions of identity between Rbx1 proteins from different species and positions of identity between the Rbx1 and APC11 proteins. Grey shading indicates positions of similarity.--

Please replace the paragraph beginning at page 9, line 13, with the following:

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--The VHL tumor suppressor complex was fractionated by 13% SDS-polyacrylamide gel electrophoresis. Proteins were visualized by staining with Coomassie blue, excised, and subjected to in-gel reduction, S-carboxyamidomethylation, and tryptic digestion. Using 10% of the digestion mixture, peptide sequences were determined in a single run by microcapillary reversed-phase chromatography coupled to the electrospray ionization source of a quadrupole ion trap mass spectrometer (Finnigan LCQ). The ion trap's online data-dependent scans allowed the automatic acquisition of high resolution spectra to determine charge state and exact mass, and tandem mass spectrometry spectra for sequence information. The relative collision energy was 35% AND ISOLATION WIDTH WAS 2.5 Dalton. Searches of the EST database performed using TLASTIN algorithm identified human and mouse ESTs that encoded the peptide sequences NHIMDLCIECQAN (SEQ ID NO:10), QVCPLDNREWEFQK (SEQ ID NO:11),

WNAVAL (SEQ ID NO:12) and WLK which were determined by ion trap mass spectrometry of the 16 kDa polypeptide that copurified with the VHL complex. The identification was facilitated with the algorithm SEQUEST (Eng, *et al.* 1994. *J Am Soc Mass Spectrom* 5:976) and by programs developed in the Harvard Microchemistry Facility (Chittum *et al.* 1998. *Biochemistry* 37:10866). I.M.A.G.E. Consortium cDNA colonies ("I.M.A.G.E. Consortium: an integrated molecular analysis of genomes and their expression," *Genomics* 33:151-152) encoding the complete 108 amino acid long ORFs of human (H71993) and mouse (W66989 and AA260889) Rbx1 were obtained from Research Genetics, Huntsville, Alabama, and the nucleotide sequences of both strands were determined. Human and mouse cDNAs encoded identical polypeptides of 108 amino acids. The amino acid sequence for human and mouse Rbx1 is shown in Fig. 2 and in SEQ ID NO:1. The nucleotide sequence for the human Rbx1 DNA is shown in nucleotides 7-333 or SEQ ID NO:3 and the nucleotide sequence for the murine Rbx1 DNA is shown in nucleotides 18-344 or SEQ ID NO:5, inclusive of the stop codon. Nucleotides 1-6 and 1017 are 5' untranslated regions, respectively and 334-508 and 345-504 are 3' untranslated regions, respectively--

Please cancel the "SEQUENCE LISTING", pages 1-2, submitted on February 25, 2000 with International Application No. PCT/US00/04838, and insert therefor the accompanying paper copy of the Substitute Sequence Listing, page numbers 1 to 5, at the end of the application.

REMARKS

The enclosed Substitute Sequence Listing corrects errors and omissions in the previously submitted Sequence Listing, filed February 25, 2000, for International Application No. PCT/US00/04838.